## This Page Is Inserted by IFW Operations and is not a part of the Official Record

## BEST AVAILABLE IMAGES

Defective images within this document are accurate representations of the original documents submitted by the applicant.

Defects in the images may include (but are not limited to):

- BLACK BORDERS
- TEXT CUT OFF AT TOP, BOTTOM OR SIDES
- FADED TEXT
- ILLEGIBLE TEXT
- SKEWED/SLANTED IMAGES
- COLORED PHOTOS
- BLACK OR VERY BLACK AND WHITE DARK PHOTOS
- GRAY SCALE DOCUMENTS

## IMAGES ARE BEST AVAILABLE COPY.

As rescanning documents will not correct images, please do not report the images to the Image Problem Mailbox.

Search results EN 00/08/1949

				ρΛ
L Number	Hits	Search Text	DB	Time stamp
2	109	Hallenbeck.in.  Hallenbeck.in. and e2f	USPAT; US-PGPUB; EPO; JPO; DERWENT; IBM_TDB USPAT;	2003/03/06 15:08 2003/03/06
			US-PGPUB; EPO; JPO; DERWENT; IBM TDB	15:08
3	2	(e2f or E2F-1 or dihyrofolate adj reductase or DHFR or DNA adj polymerase adj A or DPA or c-myc or c adj myc or B-myb or B adj myb) adj5 (promoter) same ITR and (termination or polyadenylation or "poly(A)." or poly adj A) and (adenoviral or adenovirus or viral) and cancer	USPAT; US-PGPUB; EPO; JPO; DERWENT; IBM_TDB	2003/03/06 15:26
5	173	(e2f or E2F-1 or dihyrofolate adj reductase or DHFR or DNA adj polymerase adj A or DPA or c-myc or c adj myc or B-myb or B adj myb) and (promoter) same ITR and (termination or polyadenylation or "poly(A)" or poly adj A) and (adenoviral or adenovirus)	USPAT; US-PGPUB; EPO; JPO; DERWENT; IBM_TDB	2003/03/06 15:31
4	3	(e2f or E2F-1 or dihyrofolate adj reductase or DHFR or DNA adj polymerase adj A or DPA or c-myc or c adj myc or B-myb or B adj myb) adj5 (promoter) same ITR and (termination or polyadenylation or "poly(A)" or poly adj A) and (adenoviral or adenovirus)	USPAT; US-PGPUB; EPO; JPO; DERWENT; IBM_TDB	2003/03/06 15:30
6	. 42	(e2f or E2F-1 or dihyrofolate adj reductase or DHFR or DNA adj polymerase adj A or DPA or c-myc or c adj myc or B-myb or B adj myb) and (promoter) same ITR same (termination or polyadenylation or "poly(A)" or poly adj A) and (adenoviral or adenovirus)	USPAT; US-PGPUB; EPO; JPO; DERWENT; IBM_TDB	2003/03/06 15:37
7	12	oncolytic adj5 vector	USPAT; US-PGPUB; EPO; JPO; DERWENT; IBM_TDB	2003/03/06 16:48
8	26	reductase or DHFR or DNA adj polymerase adj A or DPA or c-myc or c adj myc or B-myb or B adj myb) and (promoter) same ITR same (termination or polyadenylation or "poly(A)" or poly adj A) same (ela or elb or e4 or e2 or replication) and (adenoviral or adenovirus)	USPAT; US-PGPUB; EPO; JPO; DERWENT; IBM_TDB	2003/03/06
9	1		USPAT; US-PGPUB; EPO; JPO; DERWENT; IBM_TDB	2003/03/06 16:29
10	23	(e2f or E2F-1 or dihyrofolate adj reductase or DHFR or DNA adj polymerase adj A or DPA or c-myc or c adj myc or B-myb or B adj myb) and (promoter) same ITR same (termination or polyadenylation or "poly(A)" or poly adj A) same (e1a or e1b or e4 or e2 or replication) same (adenoviral or adenovirus)	USPAT; US-PGPUB; EPO; JPO; DERWENT; IBM_TDB	2003/03/06 16:34

Search History 3/6/03 4:50:32 PM Page 1

			T-11	10000/00/00
11	10	oncolytic and E2f	USPAT;	2003/03/06
			US-PGPUB; EPO; JPO;	16:49
			DERWENT; IBM TDB	
12	3	oncolytic and E2f and polyadenylation	USPAT;	2003/03/06
12		oncolytic and Ezi and polyadenylation	US-PGPUB;	16:49
			EPO; JPO;	
			DERWENT;	
			IBM TDB	
_	2	"10081969"	USPAT;	2003/03/06
	_		US-PGPUB;	15:08
			EPO; JPO;	
			DERWENT;	
!			IBM_TDB	
_	4255	(adenoviral or adenovirus) adj vector	USPAT;	2003/03/05
			US-PGPUB;	15:14
			EPO; JPO;	
			DERWENT;	
		[,, , , , , , , , , , , , , , , , , , ,	IBM_TDB	
_	12590	(adenoviral or adenovirus or viral) adj	USPAT;	2003/03/05
		vector	US-PGPUB;	16:19
			EPO; JPO;	
			DERWENT; IBM TDB	
_	51	oncolytic and ((adenoviral or adenovirus	USPAT;	2003/03/05
_	] 31	or viral) adj vector)	US-PGPUB;	16:20
		or virar, adj vector,	EPO; JPO;	10.20
			DERWENT;	
			IBM TDB	
_	8	e2f and (oncolytic and ((adenoviral or	USPAT;	2003/03/05
		adenovirus or viral) adj vector))	US-PGPUB;	16:20
		,,	EPO; JPO;	
			DERWENT;	
			IBM_TDB	
-	349	e2f and ((adenoviral or adenovirus or	USPAT;	2003/03/05
		viral) adj vector)	US-PGPUB;	16:20
			EPO; JPO;	
			DERWENT;	
			IBM_TDB	
-	51		USPAT;	2003/03/05
		adenovirus or viral) adj vector)	US-PGPUB;	16:21
			EPO; JPO;	
			DERWENT;	
_	1	(e2f adj5 promoter) same (termination or	IBM_TDB USPAT;	2003/03/05
_	1	(e21 adjs promoter) same (termination or   polyadenylation)	US-PGPUB;	16:26
		porgadonyracrom/	EPO; JPO;	10.20
			DERWENT;	
			IBM TDB	
_	20	(e2f or E2F-1 or dihyrofolate adj	USPAT;	2003/03/05
		reductase or DHFR or DNA adj polymerase	US-PGPUB;	16:43
		adj A or DPA or c-myc or c adj myc or	EPO; JPO;	
		B-myb or B adj myb) adj2 (promoter) same	DERWENT;	
		(termination or polyadenylation)	IBM_TDB	
-	97	(e2f or E2F-1 or dihyrofolate adj	USPAT;	2003/03/05
		reductase or DHFR or DNA adj polymerase	US-PGPUB;	16:51
		adj A or DPA or c-myc or c adj myc or	EPO; JPO;	
		B-myb or B adj myb) adj2 (promoter) and	DERWENT;	
		(termination or polyadenylation) and	IBM_TDB	
		((adenoviral or adenovirus or viral) adj		
_	1 47	vector)	IICDATE -	2002/02/05
-	147	(e2f or E2F-1 or dihyrofolate adj	USPAT;	2003/03/05
		reductase or DHFR or DNA adj polymerase adj A or DPA or c-myc or c adj myc or	US-PGPUB; EPO; JPO;	10.30
		B-myb or B adj myb) adj2 (promoter) and	DERWENT;	
		(termination or polyadenylation or	IBM TDB	
		poly(A) or poly adj A) and ((adenoviral		
		or adenovirus or viral) adj vector)		
			1	1

				1
_	51	(e2f or E2F-1 or dihyrofolate adj	USPAT;	2003/03/05
		reductase or DHFR or DNA adj polymerase	US-PGPUB;	17:10
		adj A or DPA or c-myc or c adj myc or	EPO; JPO;	
		B-myb or B adj myb) adj2 (promoter) same	DERWENT;	
		(termination or polyadenylation or	IBM_TDB	
		poly(A) or poly adj A) and ((adenoviral		
		or adenovirus or viral) adj vector)		
-	38	((e2f or E2F-1 or dihyrofolate adj	USPAT;	2003/03/06
		reductase or DHFR or DNA adj polymerase	US-PGPUB;	15:18
		adj A or DPA or c-myc or c adj myc or	EPO; JPO;	
		B-myb or B adj myb) adj2 (promoter) same	DERWENT;	
		(termination or polyadenylation or	IBM_TDB	
		poly(A) or poly adj A) and ((adenoviral	_	
		or adenovirus or viral) adj vector)) and		ł.
		cancer		
-	2	(e2f or E2F-1 or dihyrofolate adj	USPAT;	2003/03/05
		reductase or DHFR or DNA adj polymerase	US-PGPUB;	17:14
		adj A or DPA or c-myc or c adj myc or	EPO; JPO;	
		B-myb or B adj myb) adj2 (promoter) same	DERWENT;	
		(termination or polyadenylation or	IBM_TDB	i
		"poly(A)" or poly adj A) and ((adenoviral	_	
		or adenovirus or viral) adj vector)		
-	103	(e2f or E2F-1 or dihyrofolate adj	USPAT;	2003/03/05
		reductase or DHFR or DNA adj polymerase	US-PGPUB;	17:18
		adj A or DPA or c-myc or c adj myc or	EPO; JPO;	
		B-myb or B adj myb) adj2 (promoter) and	DERWENT;	
		(termination or polyadenylation or	IBM_TDB	
		"poly(A)" or poly adj A) and ((adenoviral	_	
		or adenovirus or viral) adj vector)	}	
-	73	((e2f or E2F-1 or dihyrofolate adj	USPAT;	2003/03/05
		reductase or DHFR or DNA adj polymerase	US-PGPUB;	17:18
		adj A or DPA or c-myc or c adj myc or	EPO; JPO;	
		B-myb or B adj myb) adj2 (promoter) and	DERWENT;	
		(termination or polyadenylation or	IBM_TDB	
		"poly(A)" or poly adj A) and ((adenoviral		
1		or adenovirus or viral) adj vector)) and		
		cancer		
-	1	(e2f or E2F-1 or dihyrofolate adj	USPAT;	2003/03/05
		reductase or DHFR or DNA adj polymerase	US-PGPUB;	17:21
		adj A or DPA or c-myc or c adj myc or	EPO; JPO;	
		B-myb or B adj myb) adj2 (promoter) same	DERWENT;	
		ITR and (termination or polyadenylation	IBM_TDB	
		or "poly(A)" or poly adj A) and		
		((adenoviral or adenovirus or viral) adj		
		vector)		
-	2	(e2f or E2F-1 or dihyrofolate adj	USPAT;	2003/03/05
		reductase or DHFR or DNA adj polymerase	US-PGPUB;	17:31
		adj A or DPA or c-myc or c adj myc or	EPO; JPO;	
		B-myb or B adj myb) adj2 (promoter) and	DERWENT;	
		ITR and (termination or polyadenylation	IBM_TDB	
		or "poly(A)" or poly adj A) and		
		((adenoviral or adenovirus or viral) adj		
		vector) and replication adj5 gene		

Welcome to STN International! Enter x:x LOGINID:ssspta1636mxm DASSWORD. TERMINAL (ENTER 1, 2, 3, OR ?):2 Web Page URLs for STN Seminar Schedule - N. America NEWS 2 Apr 08 "Ask CAS" for self-help around the clock NEWS 3 Apr 09 BEILSTEIN: Reload and Implementation of a New Subject Area NEWS 4 Apr 09 ZDB will be removed from STN NEWS 5 Apr 19 US Patent Applications available in IFICDB, IFIPAT, and IFIUDB NEWS 6 Apr 22 Records from IP.com available in CAPLUS, HCAPLUS, and ZCAPLUS 7 Apr 22 BIOSIS Gene Names now available in TOXCENTER NEWS 8 Apr 22 Federal Research in Progress (FEDRIP) now available NEWS 9 Jun 03 New e-mail delivery for search results now available NEWS 10 Jun 10 MEDLINE Reload NEWS 11 Jun 10 PCTFULL has been reloaded NEWS 12 Jul 02 FOREGE no longer contains STANDARDS file segment NEWS 13 Jul 22 USAN to be reloaded July 28, 2002; saved answer sets no longer valid NEWS 14 Jul 29 Enhanced polymer searching in REGISTRY NEWS 15 Jul 30 NETFIRST to be removed from STN NEWS 16 Aug 08 CANCERLIT reload NEWS 17 Aug 08 PHARMAMarketLetter(PHARMAML) - new on STN NEWS 18 Aug 08 NTIS has been reloaded and enhanced NEWS 19 Aug 19 Aquatic Toxicity Information Retrieval (AQUIRE) now available on STN NEWS 20 Aug 19 IFIPAT, IFICDB, and IFIUDB have been reloaded NEWS 21 Aug 19 The MEDLINE file segment of TOXCENTER has been reloaded NEWS 22 Aug 26 Sequence searching in REGISTRY enhanced NEWS 23 Sep 03 JAPIO has been reloaded and enhanced NEWS 24 Sep 16 Experimental properties added to the REGISTRY file NEWS 25 Sep 16 CA Section Thesaurus available in CAPLUS and CA NEWS 26 Oct 01 CASREACT Enriched with Reactions from 1907 to 1985 NEWS 27 Oct 21 EVENTLINE has been reloaded NEWS 28 Oct 24 BEILSTEIN adds new search fields NEWS 29 Oct 24 Nutraceuticals International (NUTRACEUT) now available on STN NEWS 30 Oct 25 MEDLINE SDI run of October 8, 2002 NEWS 31 Nov 18 DKILIT has been renamed APOLLIT NEWS 32 Nov 25 More calculated properties added to REGISTRY NEWS 33 Dec 02 TIBKAT will be removed from STN NEWS 34 Dec 04 CSA files on STN NEWS 35 Dec 17 PCTFULL now covers WP/PCT Applications from 1978 to date NEWS 36 Dec 17 TOXCENTER enhanced with additional content NEWS 37 Dec 17 Adis Clinical Trials Insight now available on STN NEWS 38 Dec 30 ISMEC no longer available NEWS 39 Jan 13 Indexing added to some pre-1967 records in CA/CAPLUS NEWS 40 Jan 21 NUTRACEUT offering one free connect hour in February 2003 NEWS 41 Jan 21 PHARMAML offering one free connect hour in February 2003 NEWS 42 Jan 29 Simultaneous left and right truncation added to COMPENDEX, ENERGY, INSPEC NEWS 43 Feb 13 CANCERLIT is no longer being updated NEWS 44 Feb 24 METADEX enhancements NEWS 45 Feb 24 PCTGEN now available on STN NEWS 46 Feb 24 TEMA now available on STN NEWS 47 Feb 26 NTIS now allows simultaneous left and right truncation NEWS 48 Feb 26 PCTFULL now contains images NEWS 49 Mar 04 SDI PACKAGE for monthly delivery of multifile SDI results NEWS EXPRESS January 6 CURRENT WINDOWS VERSION IS V6.01a, CURRENT MACINTOSH VERSION IS V6.0b(ENG) AND V6.0Jb(JP), AND CURRENT DISCOVER FILE IS DATED 01 OCTOBER 2002

Enter NEWS followed by the item number or name to see news on that specific topic.

General Internet Information

Welcome Banner and News Items

STN Operating Hours Plus Help Desk Availability

CAS World Wide Web Site (general information)

Direct Dial and Telecommunication Network Access to STN

NEWS HOURS

NEWS INTER

NEWS LOGIN

NEWS PHONE

NEWS WWW

All use of STN is subject to the provisions of the STN Customer agreement. Please note that this agreement limits use to scientific research. Use for software development or design or implementation of commercial gateways or other similar uses is prohibited and may result in loss of user privileges and other penalties.

FILE 'HOME' ENTERED AT 16:56:44 ON 06 MAR 2003

=> s medline caplus

THIS COMMAND NOT AVAILABLE IN THE CURRENT FILE

Some commands only work in certain files. For example, the EXPAND command can only be used to look at the index in a file which has an index. Enter "HELP COMMANDS" at an arrow prompt (=>) for a list of commands which can be used in this file.

=> file medline caplus

COST IN U.S. DOLLARS

SINCE FILE TOTAL

FULL ESTIMATED COST

**ENTRY** SESSION

0.21 0.21

FILE 'MEDLINE' ENTERED AT 16:57:02 ON 06 MAR 2003

FILE 'CAPLUS' ENTERED AT 16:57:02 ON 06 MAR 2003

USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.

PLEASE SEE "HELP USAGETERMS" FOR DETAILS.

COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)

=> s adenovirus and (promoter (A5) ITR (a5) (polyadenylation or polyA or "poly(A)" or termination)) and (e2f or DHFR or DPA or c-myc or B-myb) MISSING OPERATOR 'PROMOTER (A5'

The search profile that was entered contains terms or nested terms that are not separated by a logical operator.

=> s adenovirus and (promoter (s) ITR (s) (polyadenylation or polyA or "poly(A)" or termination)) and (e2f or DHFR or DPA or c-myc or B-myb)

2 ADENOVIRUS AND (PROMOTER (S) ITR (S) (POLYADENYLATION OR POLYA OR "POLY(A)" OR TERMINATION)) AND (E2F OR DHFR OR DPA OR C-MYC OR B-MYB)

=> d ibib abs 1-2

ANSWER 1 OF 2 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER:

2002:676177 CAPLUS

DOCUMENT NUMBER: TITLE:

137:211937 Construction of adenoviral vectors containing

insulating sequence for minimization of leaky

therapeutic gene expression

INVENTOR (S):

Gorziglia, Mario; Hallenbeck, Paul L.; Kaleko,

Michael; Clarke, Lori; Phipps, Sandrina Novartis A.-G., Switz.

PATENT ASSIGNEE(S): SOURCE:

PCT Int. Appl., 41 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATE	NT NO.		KI	ND I	DATE			A	PPLI	CATI	N NC	o. i	DATE				
								-									
WO 20	0020686	27	A	2 :	2002	0906		W	20	02-U	S528	0 :	2002	0222			
,	W: AE,	AG,	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BY,	BZ,	CA,	CH,	CN,	
	co,	CR,	CU,	CZ,	DE,	ĎK,	DM,	DZ,	EC,	EE,	ES,	FI,	GB,	GD,	GE,	GH,	
	GM,	HR,	HU,	ID,	ΙL,	IN,	IS.	JP,	KE,	KG,	KP,	KR,	KZ,	LC,	LK,	LR,	
	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MN,	MW,	MX,	MZ,	NO,	NZ,	OM,	PH,	PL,	
	PT,	RO,	RU,	SD,	SE,	SG,	SI,	SK,	SL,	TJ,	TM,	TN,	TR,	TT,	TZ,	UA,	
	UG,	US,	UZ,	VN,	YU,	ZA,	ZM.	ZW,	AM,	AZ,	BY,	KG.	KZ,	MD,	RU,	TJ,	TM
F	RW: GH,	GM,	KE,	LS,	MW,	MZ,	SD,	SL,	sz,	TZ,	UG,	ZM,	ZW,	AT,	BE,	CH,	
	CY,	DE,	DK,	ES,	FI,	FR,	GB,	GR,	IE,	IT,	LU,	MC,	NL,	PT,	SE,	TR,	
	BF,	BJ,	CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	MR,	NE,	SN,	TD,	TG	
PRIORITY A	APPLN.	INFO	. :			-	1	US 2	001-	2708	95P	P :	2001	0223			
AB The p	oresent	inv	enti	on r	elat	es to	ad	enov:	iral	vec	tors	and	the	ir u	se in	1	
metho	the	rapy	. T	he p	rese	nt i	nven	tion	pro	vide:	s no	vel '	vira	ì			
vecto	use	ful	for	he t	mini	niza	tion	of :	leak	y ger	ne e	kpre	ssion	n,			
and,	in par	ticu	lar,	of i	nons	peci	fic	tran	scri	ptio	nal:	read	thre	ough	of g	genes	3.

Such constructs may be obtained by the insertion of an insulating sequence into a vector construct, such as for example a termination signal sequence upstream of the transcription initiation site of the resp. transcription unit. Provided is a recombinant viral vector comprising an adenoviral nucleic acid backbone, wherein said nucleic acid backbone comprises in sequential order: a left ITR, a termination signal sequence, an E2F-1 promoter which is operably linked to a gene essential for replication of the recombinant viral vector, an adenoviral packaging signal, and a right ITR.

L1 ANSWER 2 OF 2 CAPLUS COPYRIGHT 2003 ACS ACCESSION NUMBER: 2002:675779 CAPLUS

DOCUMENT NUMBER: 137:210924

TITLE: Oncolytic adenoviral vectors expressing therapeutic

genes for the treatment of cancer

INVENTOR (S): Ennist, David Leonard; Forry-Schaudies, Suzanne;

> Gorziglia, Mario; Hallenbeck, Paul L.; Hay, Carl M.; Jakubczak, John Leonard; Kaleko, Michael; Ryan, Patricia Clara; Stewart, David A.; Xie, Yuefeng; Connelly, Sheila; Police, Sehidhar Reddy; Clarke,

Lori; Phipps, Sandrina; Cheng, Cheng Novartis Pharma A.-G., Switz.

PATENT ASSIGNEE(S):

SOURCE: PCT Int. Appl., 226 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

```
PATENT NO.
                   KIND DATE
                                         APPLICATION NO. DATE
                    ----
    WO 2002067861
                     A2 20020906
                                         WO 2002-US5300 20020222
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
            GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
            LS, LT, LU, LV, MA, MD, MG, MN, MW, MX, MZ, NO, NZ, OM, PH, PL,
            PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA,
            UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH,
            CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR,
            BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                                      US 2001-270922P P 20010223
PRIORITY APPLN. INFO .:
                                      US 2001-295037P P 20010601
                                      US 2002-348670P P 20020114
```

The present invention relates to oncolytic adenoviral vectors and their use in methods of gene therapy. Provided is a recombinant viral vector comprising an adenoviral nucleic acid backbone, wherein said nucleic acid backbone comprises in sequential order: a left ITR, a termination signal sequence, an E2F responsive promoter which is operably linked to a gene essential for replication of the recombinant viral vector, an adenoviral packaging signal, and a right ITR. The adenoviral vectors may also comprise a polynucleotide encoding a cytokine such as GM-CSF that can stimulate a systemic immune response against tumor cells. The preferred vector Ar6pAE2fF comprises an adenovirus vector that uses a fragment of the human E2F-1 promoter to selectively regulate E1A expression and thus adenoviral replication in tumor cells. Ar6pAE2fF selectively kills Rb-pathway defective tumor cells over normal primary cells, and is preferentially replicated in human tumor cell lines vs. normal primary cells. This vector has a superior early toxicity profile to the non-selective replication competent virus, Addl327, when administered i.v. in SCID mice and provides advantages in efficacy, selectivity, and safety as compared to the oncolytic viral vector Addl1520. Ar17pAE2fTrtex is a particularly preferred, tumor-selective oncolytic adenovirus designed for the treatment of a broad range of cancer indications involving the two most common alterations in human cancer, namely defects in the Rb-pathway and overexpression of telomerase. Ar17pAE2fTrtex utilizes a E2F-1 promoter to control expression of the adenoviral E1A gene and the adenoviral E4 gene is controlled by a hTERT (human telomerase reverse transcriptase) promoter. Ar17pAE2fTrtex is expected to replicate in the majority of cancer cells, lead to tumor selective expression of toxic viral proteins, cytolysis, and enhancement of sensitivity to chemotherapy, cytokines, and cytotoxic T lymphocytes.

=> FIL STNGUIDE COST IN U.S. DOLLARS

ENTRY SESSION FULL ESTIMATED COST 32.67 32.88

SINCE FILE

TOTAL

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

SINCE FILE
ENTRY
SESSION

CA SUBSCRIBER PRICE

-1.30
-1.30

FILE 'STNGUIDE' ENTERED AT 17:00:27 ON 06 MAR 2003
USE IS SUBJECT TO THE TERMS OF YOUR CUSTOMER AGREEMENT
COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY, JAPAN SCIENCE
AND TECHNOLOGY CORPORATION, AND FACHINFORMATIONSZENTRUM KARLSRUHE

FILE CONTAINS CURRENT INFORMATION.
LAST RELOADED: Feb 28, 2003 (20030228/UP).

=> file medline caplus

 COST IN U.S. DOLLARS
 SINCE FILE TOTAL

 ENTRY
 SESSION

 FULL ESTIMATED COST
 0.06
 32.94

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

SINCE FILE TOTAL
ENTRY SESSION
CA SUBSCRIBER PRICE

0.00 -1.30

FILE 'MEDLINE' ENTERED AT 17:01:16 ON 06 MAR 2003

FILE 'CAPLUS' ENTERED AT 17:01:16 ON 06 MAR 2003 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)

=> d his

(FILE 'HOME' ENTERED AT 16:56:44 ON 06 MAR 2003)

FILE 'MEDLINE, CAPLUS' ENTERED AT 16:57:02 ON 06 MAR 2003
L1 2 S ADENOVIRUS AND (PROMOTER (S) ITR (S) (POLYADENYLATION OR POLY

FILE 'STNGUIDE' ENTERED AT 17:00:27 ON 06 MAR 2003

FILE 'MEDLINE, CAPLUS' ENTERED AT 17:01:16 ON 06 MAR 2003

=> s adenovirus and (promoter (s) ITR (s) (polyadenylation or polyA or "poly(A)" or termination))
L2 8 ADENOVIRUS AND (PROMOTER (S) ITR (S) (POLYADENYLATION OR POLYA
OR "POLY(A)" OR TERMINATION))

=> dup remove 12

PROCESSING COMPLETED FOR L2

L3 7 DUP REMOVE L2 (1 DUPLICATE REMOVED)

=> s 13 and py<=2001\

NUMERIC VALUE NOT VALID '2001\'

NUMERIC VALUE NOT VALID '2001\'

Numeric values may contain 1-8 significant figures. If range notation is used, both the beginning and the end of the range must be specified, e.g., '250-300/MW'. Expressions such as '250-/MW' are not allowed. To search for values above or below a given number, use the >, =>, <, or <= operators, e.g., 'MW => 250'. Text terms cannot be used in numeric expressions. If you specify a unit, it must be dimensionally correct for that field code. To see the unit designations for field codes in the current file, enter "DISPLAY UNIT ALL" at an arrow prompt (=>).

=> s 13 and py<=2001

L4 5 L3 AND PY<=2001

=> d ibib abs 1-5

L4 ANSWER 1 OF 5 MEDLINE

ACCESSION NUMBER: 1999373450 MEDLINE

DOCUMENT NUMBER: 99373450 PubMed ID: 10441562

TITLE: Transcription map and expression of bovine herpesvirus-1

glycoprotein D in early region 4 of bovine

adenovirus-3.

AUTHOR: Baxi M K; Babiuk L A; Mehtali M; Tikoo S K

CORPORATE SOURCE: Veterinary Infectious Disease Organization, University of

Saskatchewan, Saskatoon, Saskatchewan, S7N 5E3, Canada.

SOURCE: VIROLOGY, (1999 Aug 15) 261 (1) 143-52.

Journal code: 0110674. ISSN: 0042-6822.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199909

ENTRY DATE: Entered STN: 19990925

Last Updated on STN: 19990925 Entered Medline: 19990907

AB Early region 4 (E4) of bovine adenovirus type 3 (BAV-3) was analyzed by Northern blotting, RT-PCR analysis, cDNA sequencing, and S1 nuclease protection assays. The transcriptional map of the E4 region of BAV-3 has marked dissimilarities from those of mouse adenovirus -1, ovine adenovirus-287, and human adenovirus-2, for which the transcriptional maps have been constructed. The E4 region of

which the transcriptional maps have been constructed. The E4 region of BAV-3, located between 98.6 and 89.8 MU transcribes seven distinct classes of bovine adenovirus type 3 mRNA. The seven mRNA species formed by the removal of one to three introns share both the 3' end and a short 5' leader (25 nucleotides). The E4 mRNAs can encode at least five unique polypeptides, namely, 143R1, 69R, 143R2, 268R, and 219R. Isolation of a replication-competent recombinant "BAV404" containing 1.9-kb insertion [glycoprotein (gD) of bovine herpesvirus 1, under the control of a SV40 early promoter and poly(A)] in the region

between E4 and the right ITR suggested that this region is nonessential for BAV-3 replication. Expression of gD by BAV404 recombinant virus was confirmed by immunoprecipitation with gD-specific monoclonal antibodies. Analysis of the kinetics of protein expression indicated that gD is expressed at both early and late times postinfection. These results suggest that: (a) E4 produces seven 5'-3' coterminal mRNAs and (b) the right terminal region of BAV-3 can be used for the expression of vaccine anticens.

Copyright 1999 Academic Press.

L4 ANSWER 2 OF 5 MEDLINE

ACCESSION NUMBER: 90223999 MEDLINE

DOCUMENT NUMBER: 90223999 PubMed ID: 2183470

TITLE: High level expression of the envelope glycoproteins of the

human immunodeficiency virus type I in presence of rev gene

using helper-independent adenovirus type 7

recombinants.

AUTHOR: Chanda P K; Natuk R J; Mason B B; Bhat B M; Greenberg L;

Dheer S K; Molnar-Kimber K L; Mizutani S; Lubeck M D; Davis

A R; +

CORPORATE SOURCE: Biotechnology and Microbiology Division, Wyeth-Ayerst

Research, Philadelphia, Pennsylvania 19101.
SOURCE: VIROLOGY, (1990 Apr) 175 (2) 535-47.

Journal code: 0110674. ISSN: 0042-6822.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals; AIDS

ENTRY MONTH: 199005

ENTRY DATE: Entered STN: 19900622

Last Updated on STN: 19970203 Entered Medline: 19900523

AB The effect of rev (art/trs) gene on the level of HIV-1 envelope (env) expression using recombinant adenovirus was investigated. Recombinant adenoviruses expressing either the envelope or the rev gene of the human immunodeficiency virus type 1 (HIV-1) were constructed by inserting the gene into an expression cassette. The expression cassette contained the adenovirus type 7 major late promoter, followed by leader 1 of the adenovirus tripartite leader and a portion of intron between leaders 1 and 2, le 2 and 3, and a hexon polyadenylation signal. The cassette was

tripartite leader and a portion of intron between leaders 1 and 2, leaders then inserted at the terminal region between the E4 and ITR regions of the adenovirus 7 genome with a concomitant E3 region deletion (80-87 m.u.). A549 cells infected with the recombinant virus containing the env gene produced the envelope glycoproteins gp160, gp120, and gp41. HIV-1 envelope gene expression was greatly enhanced (20- to 50-fold) in the cells that were simultaneously infected with the recombinant adenovirus containing the rev gene as measured by ELISA and Western blotting. Interestingly, this effect was observed despite the lack of the 5' down splice site for rev and seems to be post-transcriptional. Another recombinant adenovirus which contains both the rev and the env genes was constructed by inserting the rev gene in the deleted E3 region and the env gene in the terminal cassette. This double recombinant virus expressed high levels of env antigen in A549 cells similar to those attained upon co-infection with two separate recombinant viruses containing the rev or env gene. Furthermore, the rev gene nucleotide sequence could be altered without altering the amino acid sequence and its sequences truncated by 17 amino acids from the C-terminus had no effect of rev function.

DOCUMENT NUMBER:

130:135015

TITLE:

Cloning vectors for producing adenoviral minimal

viruses

INVENTOR (S):

Hillgenberg, Moritz; Loser, Peter; Schnieders, Frank;

Sandig, Volker; Strauss, Michael

PATENT ASSIGNEE(S):

Hepavec A.-G. fur Gentherapie, Germany

SOURCE:

PR I

PCT Int. Appl., 57 pp.

DOCUMENT TYPE:

CODEN: PIXXD2

Patent

LANGUAGE:

German

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

	PAT	rent	NO.		KI	ND	DATE			Α	PPLI	CATI	ON NO	٥.	DATE			
		- <b></b> .								-								
	WO	9902	2647		A:	2	1999	0121		W	0 19	98-D	E194	0	1998	0706	<	
	WO	9902	647		A:	3	1999	0415										
		W:	JP,	US														
		RW:	AT,	BE,	CH,	CY	, DE,	DK,	ES,	FI,	FR,	GB,	GR,	IE,	, IT,	LU,	MC,	NL.
			PT,	SE														
	DE	1974	4768	3	A	1	2000	0803		D	E 19	97-1	9744	768	1997	1010	<	
	DE	1974	4768	3	C	2	2002	0411		•								
	ΕP	1003	895		A:	2	2000	0531		E	P 19	98-9	4498	4	1998	0706	<	
		R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	ΙT,	LI,	NL,	SE				
	JP	2001	15093	375	T	2	2001	0724		J	P 20	00-5	0214	3	1998	0706	<	
0	RITY	API	LN.	INFO	. :				1	DE 1	997-	1972	9571	Α	1997	0710		
									1	DE 1	997-	1974	4768	Α	1997	1010		
									1	WO 1	998-	DE19	40	W	1998	0706		

AB The invention relates to cloning vectors for producing adenoviral minimal viruses, consisting of: a) two adenoviral inverted terminal repeats (ITRs) which are flanked by ab) two restriction sites with a recognition sequence more than 8 bp in length, and enclose ac) an adenoviral packaging signal, ad) a multiple cloning site for inserting therapeutic DNA fragments into which non-coding mammalian chromosomal DNA may addnl. be cloned, ae) (optionally) a recognition site for a recombinase situated between one of the ITRs and the adenoviral packaging signal, and af) (optionally) a reporter gene cassette; b) a bacterial plasmid backbone with replication origin and bacterial resistance gene, into which ba) a packaging signal of a bacteriophage is cloned. Thus, minimal adenoviral vector cloning plasmid pMVX-Bg was created. This plasmid consists of an adenovirus 5 5'-ITR, a human chromosomal stuffer, a Rouse sarcoma virus promoter linked to a lacZ gene and an SV40 polyA sequence, an adenovirus 5 3'-ITR, an I-SceI restriction site, an ampR gene, a cos site, and another I-SceI

ANSWER 4 OF 5 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER:

restriction site.

1998:605015 CAPLUS

DOCUMENT NUMBER:

129:198915

TITLE:

Expression vector for the permanent expression of

foreign DNA

INVENTOR (S):

Grummt, Ingrid; Grummt, Friedrich

PATENT ASSIGNEE(S):

Deutsches Krebsforschungszentrum Stiftung des Offentlichen Rechts, Germany

PCT Int. Appl., 10 pp.

SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE:

Patent German

LANGUAGE:

FAMILY ACC. NUM. COUNT: 1 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9837209	A2	19980827	WO 1998-DE539	19980224 <
WO 9837209	A3	19981126		
W: CA,	JP, US			
RW: AT,	BE, CH, DE	, DK, ES, F	I, FR, GB, GR, IE, IT,	, LU, MC, NL, PT, SE
DE 19707273	C1	19980924	DE 1997-19707273	19970224 <
EP 968296	A2	20000105	EP 1998-914811	19980224 <
R: AT,	BE, CH, DE	, DK, ES, FI	R, GB, IT, LI, NL, SE	
JP 20015123	20 T2	20010821	JP 1998-536164	19980224 <
US 6300126	B1	20011009	US 1999-367927	19991020 <
PRIORITY APPLN.	INFO.:		DE 1997-19707273 A	19970224
			WO 1998-DE539 W	19980224

The present invention relates to an expression vector for expressing foreign DNA. Said DNA at its 3' end has a sequence which prevents the replication of the expression vector from occurring in the opposite direction to the transcription of said expression vector. The invention also relates to a prepn. contg. such an expression vector and to the use of both in the permanent expression of foreign DNA in cells. Thus,

expression vector pAAV-ADA, comprising adeno-assocd. virus 5'- and 3'-ITRs, mouse metallothionein promoter, human adenosine deaminase cDNA, SV40 poly A sequence, and a replication fork barrier, was prepd. COS cells infected with adenovirus and expressing AAV rep and cap genes were used to prep. virus particles. Infection of cells with these virus particles led to permanent expression of the ADA gene.

```
ANSWER 5 OF 5 CAPLUS COPYRIGHT 2003 ACS
ACCESSION NUMBER:
                        1995:324646 CAPLUS
DOCUMENT NUMBER:
                         122:153378
TITLE:
                         Adenoviral vectors containing DNA encoding human lung
                         surfactant protein
INVENTOR (S):
                        Trapnell, Bruce; Whitsett, Jeffrey
                        Genetic Therapy, Inc., USA; University of Cincinnati
PATENT ASSIGNEE(S):
                         PCT Int. Appl., 42 pp.
SOURCE:
                         CODEN: PIXXD2
DOCUMENT TYPE:
                         Patent
LANGUAGE:
                         English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:
    PATENT NO.
                     KIND DATE
                                          APPLICATION NO. DATE
```

WO 9423582 19941027 WO 1994-US3831 19940407 <--A1 W: CA, JP, US RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE CA 2160136 . AA 19941027 CA 1994-2160136 19940407 <--A1 19960320 EP 1994-914075 19940407 <--R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE JP 09500782 JP 1994-523310 19940407 <--T2 19970128 PRIORITY APPLN. INFO.: US 1993-44406 19930408 WO 1994-US3831 19940407

Adenoviral vectors contg. a DNA sequence encoding a lung surfactant protein are described. The adenoviral vector may be a replication-deficient adenoviral vector which is free of at least the majority of the E1 and E3 DNA sequences. Thus, the recombinant adenoviral vector AV1SPB1 contg. human surfactant protein B (SPB) cDNA was constructed through homologous recombination between the adenovirus 5 deletion mutant Ad-d1327 and plasmid pAVS6SPB#7. Ad-d1327 has a deleted E3 region in which base pairs 28,593-30,470 are absent. Plasmid pAVS6SPB#7 contains an adenoviral 5'-ITR, an origin of replication contained completely within the 5'-ITR, an Ela enhancer and encapsidation signal, a Rous sarcoma virus promoter, and adenovirus 5' tripartite leader sequence, and the 2-kb human SPB cDNA including the entire protein coding sequence (nucleotides 1-1172), and the SV40 poly(A) signal. Such vectors may be employed for generation of infectious viral particles which may transduce lung epithelial cells in vivo to enable the expression of surfactant protein by such cells. The adenoviral vectors can treat lung surfactant protein deficiency states such as infant respiratory distress syndrome or adult respiratory distress syndrome.

=>

Connection closed by remote host